

(19) World Intellectual Property
Organization
International Bureau



(43) International Publication Date
6 January 2005 (06.01.2005)

PCT

(10) International Publication Number
WO 2005/001053 A3

(51) International Patent Classification⁷: A61K 39/385,
39/395

GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE,
KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD,
MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG,
PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM,
TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM,
ZW.

(21) International Application Number:
PCT/US2004/018451

(22) International Filing Date: 9 June 2004 (09.06.2004)

(84) Designated States (unless otherwise indicated, for every
kind of regional protection available): ARIPO (BW, GH,
GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM,
ZW), Eurasian (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM),
European (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI,
FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI,
SK, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ,
GW, ML, MR, NE, SN, TD, TG).

(25) Filing Language: English

(26) Publication Language: English

(30) Priority Data:
60/477,796 9 June 2003 (09.06.2003) US

Published:

- with international search report
- before the expiration of the time limit for amending the claims and to be republished in the event of receipt of amendments

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(88) Date of publication of the international search report:
11 August 2005

(81) Designated States (unless otherwise indicated, for every
kind of national protection available): AE, AG, AL, AM,
AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN,
CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI,

For two-letter codes and other abbreviations, refer to the "Guid-
ance Notes on Codes and Abbreviations" appearing at the begin-
ning of each regular issue of the PCT Gazette.

A3

WO 2005/001053

(54) Title: METHOD OF INHIBITING RECEPTOR TYROSINE KINASES WITH AN EXTRACELLULAR ANTAGONIST
AND AN INTRACELLULAR ANTAGONIST

(57) Abstract: The present invention relates to methods of inhibiting receptor tyrosine kinases by utilizing a combination of both an extracellular and an intracellular RTK antagonist. The extracellular RTK antagonist is a biological molecule or a small molecule that inhibits activation of the receptor tyrosine kinase by interacting with the extracellular binding region of the receptor. The intracellular RTK antagonist is a biological molecule or small molecule that inhibits tyrosine kinase activity of the receptor tyrosine kinase by interacting with the receptor's intracellular region bearing a kinase domain or by interacting with an intracellular protein involved in the signaling pathway of the receptor tyrosine kinase. The present invention also provides methods of treating tyrosine kinase-dependent diseases, and compositions for use in such methods thereof, by administering a combination of both an extracellular and an intracellular RTK antagonist.